Therapeutic Possibilities of Stem Cell Research

Heather Payne Genomics, Bioinformatics and Medicine Doug Brutlag May 19, 2005



http://www.stemcellresearchfoundation.org/WhatsNew/EmbryonicStemCells.htm



http://www.stemcellresearchfoundation.org/WhatsNew/PSA_1.htm http://www.stemcellresearchfoundation.org/WhatsNew/PSA_2.htm

Olfactory Bulb Stem Cells

- Primitive stem cells that normally feed the constant, life-long regeneration of odor-detecting nerves
- Like embryonic stem cells, they develop into many different types of cells in the right chemical or cellular environment
- Fairly accessible, readily obtained in all individuals and easy to grow and multiply
- Potential non-embryonic source for cells that could prove useful in replacing nerve cells lost due to injury or diseases like ALS and Parkinson's
- Transplant not subject to immune rejection

Lou Gehrig's Disease (ALS)

- Mouse experiments
 Neuronal stem cells transplanted into spinal cord
 Significantly prolonged lives by becoming neurons and interacting with existing neurons
 Symptoms developed at 137 days verses 90 days
- Treated mice lived 2 months longer



A neuromuscular disease that attacks the nerve cells and pathways in the brain and spinal cord



Heart Disease



Cause of Death Rank: #1 (2002) Morbidity: 23 million diagnosed adults (2002)Hospital Inpatient Care: 4.4 million discharges, 4.6 day length of stay (2002) Hospice Care: 13,500 (2000) Nursing Home Residents: 165,100 (1999) Morality: 696,947 (2002)

Heart Disease

- University of Pittsburgh Medical Center
- Treatment for congestive heart failure and other heart disease
- CD34+ bone marrow-derived stem cells
- Cells and blood plasma injected into 25 to 30 sites of the diseased heart (25-45 m. cells)
- New heart cells and blood vessels
- Do stem cells take on the functional characteristics of heart cells and blood vessels, or do they recruit other cells and growth factors to help regenerate heart tissue?

Bone Repair

- NJ Institute of Technology use of stem cells to induce bone repair
- Adult Stem Cells mixed with biomaterials known as scaffolds to regenerate bone growth
- Stem Cells from one person can successfully implant in another
- Diabetes, osteoporosis, cancer surgeries
- Also testing biomaterials that may repair cartilage, tendons and neuronal tissue

Type I Diabetes

- Used protein transduction to promote pancreatic cell differentiation from adult bone marrow stem cells
- Enables development of more insulin-producing cells, or islets, for transplantation into patients with Type I Diabetes
- Know the genes to make a stem cell into an insulinproducing pancreatic cell, and found the signals involved in their activation
- Can turn genes on in the right order to get functioning cells
- Can add and remove proteins as needed from the developmental process which may give advantages over less flexible approaches such as gene therapy
- Bone marrow-derived stem cells may not have the same antigen as pancreatic beta cells, which would eliminate the potential for rejection or a negative immune response

Spinal Cord Injury

Rat Experiments Sensory and motor deficiencies; paralysis Treatment derived from human embryonic stem cells and must occur in the acute phase of spinal stabilization Cells differentiate into earlystage oligodendrocytes, the building blocks of myelin Transplanted cells migrated to appropriate neuronal sites in the spinal cord 7 days post injury vs. 10 months post injury



DEBBIE HILL/UPI

Cystic Fibrosis: Stem Cell-Gene Therapy Approach

- Human bone marrowderived stem cells can differentiate into airway epithelial cells
- Encoding these cells with the gene that is defective in CF restores cellular function
- Keep airways clear of mucus and air-borne irritants
- Hope to perform clinical trial in next 2-3 years





Biological Pacemaker



- Human Embryonic Stem Cells genetically engineered and coaxed to become heart cells
- Clusters of cells beat on their own triggered the unified beating of rat heart muscle cells
- Triggered regular beating when implanted in guinea pigs
- Cells responded to drugs used to slow or speed up heart rate
- Use genetic engineering to customize the pacing rate of the cells



Hemophilia



- UNC Chapel Hill Medical School treated embryonic stem cells with fibroblast growth factor for seven days prior to injection Differentiate into early endoderm precursors Engraft, persist, differentiate further and function following injection - resulting in persistent production of Factor IX (hepatocytes) Robustly engrafted in the liver and not recognized as foreign by immune system Cells became hepatocytes 4 months later mice still producing Factor IX without immune rejection or suppression
- Low incidence of teratoma

Hearing Impairment

 Indiana University School of Medicine transformed adult bone marrow stem cells into cells with many characteristics of sensory nerve cells found in the ears

 Marrow-stromal cells develop into fat, bone and cartilage

 Autologous cell-based therapy to stimulate growth of nerve cells often missing in the inner ear of patients with profound hearing loss

Tooth Replacements

 Adult stem cells harvested from baby or wisdom teeth to grow new teeth naturally

 Contain rich supplies of stem cells that can develop into a variety of cell types including tooth generating cells

Gingivitis and periodontis



Cosmetic and Reconstructive Surgery

- Conventional soft tissue implants lose 40 to 60% of volume
- Stem cell generated natural tissues instead of synthetic implants
- Avoid problems of saline and silicon
- Won't shrink or lose shape
- Mouse experiments: bone marrow stem cells placed under the skin for four weeks; stem cells differentiated into fat generating cells and implants retained original size and shape
- Breast cancer surgery, post-cancer facial soft tissue reconstruction, trauma surgeries

Retinal Degeneration

- Mice predisposed for Retinitis Pigmentosa: a degenerative disease that destroys retinas
- Injected bone marrowderived stem cells into the back of mouse eyes during development
- Dramatically curtailed retinal degeneration
- Completely normal vasculature, improved retinal tissue and light response
- Disorders of the retina that have vascular and neuronal degeneration: genetic disorders known collectively as retinitis pigmentosa



Fig. 6. Vertical sagittal section of the adult human eye.

Potential of Embryonic Stem Cells

Research Promise: Fundamental new tools to study everything from cancer to Alzheimer's

Medical Promise: Therapies that might eventually treat dozens of diseases

Potential of Embryonic Stem Cells

- Cancer therapy: stem cells to replace organ tissue or regrow organs
- Transplants for autoimmune disorders like Lupus
- Organ transplants without risk of rejection
- Grow insulin producing Beta cells to treat Type I Diabetes
- Neurons to restore brain function to patients with Parkinson's Disease
- Rebuild an injured spine
- Create replacement skin for a burn victim
- Alzheimer's Disease, Lou Gehrig's Disease, "Bubble Children"

Opposition

- Human embryo moral status
- Opens the door to reproductive cloning - implantation to difficult to enforce
- Perfecting nuclear transplantation techniques that can be used to clone humans
- Destruction of innocent human life for research
- Biotechnical abolition of the human family
- Degradation of human procreation, manufacturing children without normal familial bonds, turning pregnancy into a research



Weldon-Stupak Bill & Brownback Bill

